### NAVAL HEALTH RESEARCH CENTER

# GULF WAR VETERAN HEALTH CARE UTILIZATION: AFTER MORE THAN 10 YEARS OF STUDY, WHAT HAVE WE LEARNED?

G. C. Gray
G. D. Gackstetter
H. K. Kang
J. T. Graham
K. C. Scott

20040121 041

Report No. 00-26

Approved for public release; distribution unlimited.

NAVAL HEALTH RESEARCH CENTER P. O. BOX 85122 SAN DIEGO, CA 92186-5122

W.

BUREAU OF MEDICINE AND SURGERY (MED-02) 2300 E ST. NW WASHINGTON, DC 20372-5300



## Gulf War Veteran Healthcare Utilization: After More than 10 Years of Study, What Have We Learned?

Gregory C. Gray,<sup>1,6</sup>
Gary D. Gackstetter,<sup>2</sup>
Han K. Kang,<sup>3</sup>
John T. Graham,<sup>4</sup>
Ken C. Scott<sup>5</sup>

<sup>1</sup> Department of Epidemiology College of Public Health University of Iowa, Iowa City, IA

<sup>2</sup>The Uniformed Services University of the Health Sciences Bethesda, MD

<sup>3</sup>The Environmental Epidemiology Service Department of Veterans Affairs Washington, DC

<sup>4</sup>British Liaison Officer (Gulf Health)
Defence Staff, British Embassy
Washington, DC

<sup>5</sup>Director of Medical Policy Canadian Forces, Ottawa Ontario, Canada

<sup>6</sup>Naval Health Research Center San Diego, CA

April 25, 2003

Report 00-26, supported by the Office of the Assistant Secretary of Defense, Health Affairs, under DoD/HA reimbursable funds work unit no. 60002. The views expressed in this article are those of the authors and do not reflect the official policy or position of the Department of the Defense, Department of Veterans Affairs, the US Government, the Ministry of Defence or the British Government. Approved for public release, distribution unlimited. This review has been conducted in compliance with all applicable Federal Regulations governing the protection of human subjects in research.

#### INTRODUCTION

Soon after returning from the Gulf War conflict, veterans from the United States, United Kingdom, and Canada became concerned that their symptoms and illnesses were due to military service. (1-3) After two United States (US) epidemic investigations (4.5) were unable to construct a case definition of a war-related syndrome or to implicate specific wartime exposures, special healthcare registries were established to systematically evaluate the health of Gulf War veterans (Table 1). As of June 2002, more than 120,000 US, United Kingdom (UK), and Canadian Gulf War veterans have been systematically evaluated under these clinical programs. In conjunction with the health registry evaluations, the US, UK, and Canada commissioned numerous medical research projects to examine whether postwar morbidity was associated with Gulf War service. (10, 11) Additional investigative efforts have been made to understand these illnesses through numerous external panel reviews and a large risk management organization. It has been estimated that in total more than 1 billion dollars has been invested in understanding and treating Gulf War veterans' illnesses (COL James R. Riddle, USAF, Executive Secretary of the Armed Forces Epidemiological Board).

We sought to summarize what has been learned from these efforts. As other authors have recently reviewed various studies of Gulf War veterans' health including mortality studies, (12) symptom studies, infectious diseases reports, and healthcare policy changes, we narrowed our review to studies involving Gulf War veteran healthcare utilization.

#### **METHODS**

We used MEDLINE, Expert Panel Reports, Reports to Congress, and a topical Gulf War veteran bibliography with 5520 references<sup>(17)</sup> to identify published research, and ongoing research relevant to the objectives. We summarize their findings herein.

#### RESULTS

Gulf War veteran healthcare registry studies

A number of research teams have evaluated Gulf War veteran healthcare registry data. (7,8,18-21) Because of volunteer bias, sensational media reports, recall bias, and other potential confounders of self-reported information, (18, 20, 22) these data are limited in their epidemiological value. However, the registry data have been very useful in documenting both the wide spectrum of self-reported symptoms and the most commonly recorded diagnoses. Additionally, the healthcare registry programs have improved Gulf War veteran access to medical care.

The methodology and findings of the US Department of Veterans Affairs (VA) and Department of Defense (DoD) healthcare registry evaluation processes have been

previously summarized in detail.<sup>(7, 19)</sup> In brief, enrollment for registry evaluation is gained by directly contacting nearby VA or DoD medical facilities or via toll-free telephone calls for appointment. Participants normally receive an initial screening examination by a primary care healthcare provider. Veterans also complete a questionnaire that enumerates medical conditions, symptoms, and possible exposures. After the initial evaluation, some Gulf War veterans may be referred for additional diagnostic procedures involving clinician-directed screenings for various immunologic, metabolic, rheumatogenic, infectious, and mental health conditions.

The UK medical assessment program<sup>(8)</sup> similarly offers veterans and their partners an evaluation by a consultant physician. Before the appointment, available service medical records are retrieved. During the consultation, patients complete a short questionnaire about their military service and relevant experiences in the Gulf, including exposure to potentially harmful factors. A detailed history is then obtained, and a clinical examination is conducted to include a complete blood count, blood chemistry tests, serology for a number of infectious agents, urinalysis, electrocar-diography, chest radiography, abdominal ultrasonography, and lung function tests. Further investigations are conducted if clinically indicated. The results of the clinical examination and investigations are sent to the patient's general practitioner. Referral to a specialist, if indicated, is either arranged within the Defence Medical Services or, if the veteran has left military service, recommended to the patient's general practitioner. <sup>(8)</sup>

The Canadian Gulf War Registry was established in January 1995. Veterans with or without medical problems were evaluated at the local base level. If a diagnosis was reached and the member and examining physician were satisfied, no further workup or referral was done. However, if the member or examining physician requested further assistance, a referral could be made to a single Gulf War Clinic located in Ottawa, Ontario. At the Gulf War Clinic, 104 veterans were evaluated, with comprehensive examinations and laboratory testing lasting on average 3 hours. The Canadian Gulf War Clinic closed in December 1997. After seeing 226 veterans, the Gulf War Registry ceased recording new entries as of January 1998. Since then, approximately 20 additional veterans have been evaluated by multiple Post-Deployment Clinics across Canada.

The first peer-reviewed Gulf War registry clinical series was published by DoD medical personnel in early 1997.<sup>(7)</sup> The authors reported a wide range of symptom reporting and clinical diagnoses, concluding that no predominant illness or suggestion of a new syndrome could be found. Their summary of evaluations conducted at 184 different medical treatment facilities was important in demonstrating the prevalence of the types of commonly reported symptoms and commonly diagnosed conditions among those who chose to be evaluated.

In 1998, DoD and VA investigators published a risk factor analysis of participation in either the DoD or VA healthcare registries. (18) Evaluating 74,653 registry participants, the study demonstrated that Gulf War veterans who were Army personnel, National

Guard and Reserve personnel, older Gulf War veterans, and personnel hospitalized before the war were most likely to seek healthcare evaluations. About 28% of registry participants were not given a specific diagnoses. The authors also identified a possible association between registry participation and news stories involving Gulf War veterans.

A summary of the United Kingdom's findings with its first 1000 registry evaluations was published in 1999. (8) Many Gulf war veterans had a wide variety of symptoms. This initial review showed no evidence of a single illness, psychological or physical, to explain the pattern of symptoms seen in veterans in the assessment program. More than one half of the veterans had more than one diagnosed condition. One third had at least one condition for which no firm somatic or psychological diagnosis could be given. In approximately 10% of veterans, no other main diagnosis was made. Conditions characterized by fatigue were found in one quarter of patients. Post-traumatic stress disorder was diagnosed in approximately 10% of patients.

The largest number of Gulf War veteran registry evaluations has been conducted at VA medical treatment facilities. A 1999 review of data from 52,835 VA registry participants listed a wide variety of symptoms and diagnoses without an apparent internal variation by military characteristics, such as branch of service (Army, Navy, Air Force, Marines) and service component (active, Reserve, National Guard). (19) The frequency of major self-reported symptoms (fatigue, skin rash, headache, muscle and joint pain, and memory loss) showed an increasing trend over a 2.5-year evaluation period (August 1992 to July 1995), but the proportion of veterans who received a physician's diagnosis of any medical condition remained fairly constant at about 72% to 76% during the same period. No single category of disease increased or decreased substantially over time. Among these veteran participants, the most frequent complaints included fatigue, skin rash, headache, muscle and joint pain, cognitive complaints, shortness of breath, sleep disturbances, diarrhea and other gastrointestinal symptoms, A significant proportion of veterans had no health complaints at the and chest pain. time of their registry examination and just wanted to participate. Within the broad diagnostic categories, the most frequently diagnosed medical conditions were diseases of musculoskeletal and connective tissue, followed by mental diseases, diseases of the respiratory system, skin and subcutaneous tissue diseases, and digestive diseases.

Reviewing existing data and reports, the most frequently reported symptoms and diagnoses from the four registries were remarkably similar. Fatigue, rashes, headache, muscle and joint pain, and cognitive problems were all quite common (Table 2), as were related, often nonspecific, diagnoses (Table 3). These findings have been important in consistently demonstrating the broad spectrum of symptoms and illnesses Gulf War veterans have experienced and their similarity to the clinical findings from other nonmilitary adult outpatient populations. (23)

Gulf War veteran registries have also been used to examine self-reported symptom and exposure data for possible illness associations and to evaluate the etiological nature of increased symptom reporting. In a June 1998 report, Kroenke et al. (23)

examined records from 18,495 DoD evaluations and concluded that the latency of symptom reporting (often more than 1 year after the war) and the poor correlation with self-reported war exposures did not support the hypothesis that a specific exposure(s) resulted in postwar symptoms. In a 1998 case series, Escalante and Fischback<sup>(24)</sup> tabulated the rheumatic manifestations of 145 VA registry participants who were referred for rheumatological evaluation. The authors noted that the types of conditions Gulf War veterans experienced after the war were not unusual (e.g. fibromvalgia. nonspecific arthralgias, osteoarthritis) and they inferred no suggestion of a new Gulf War-related syndrome. Two other teams of researchers (25, 26) studied a total of 457 Gulf War veterans evaluated for possible rheumatological conditions and noted a high prevalence of common conditions, also suggesting no unique Gulf War diagnosis or condition. Among the first 65 DoD registry patients seen for neurological symptoms at one medical treatment facility during 1994, Newmark et al. (27) found no consistent patterns of neurologic disease. In 1998, Roy et al. (28) studied registry data from 12,744 DoD registry participants who had principal or secondary diagnoses coded in the "signs, symptoms, and illdefined conditions" diagnostic category of the International Classification of Diseases, 9th revision (ICD-9). They noted that many of the diagnoses in this category represented common symptoms and concluded that there was no evidence of a new syndrome. In a 1999 report, Kipen et al. (30) surveyed 1935 VA registry participants and found that 16% of registry participants reported vague symptoms consistent with chronic fatigue syndrome, further suggesting no evidence of a new disease entity.

In contrast to the above-mentioned reports of no unusual morbidity, Das et al. (31) performed pulmonary function testing (with bronchoscopy studies on 17) on a case series of 48 VA Gulf War veteran participants referred for pulmonary medicine evaluation. The authors reported a higher than expected midvital flow capacity with the suggestion of chronic inflammation of upper airways. They speculated that these findings may be chronic manifestations of exposure to smoke and/or other irritants from the Kuwaiti oil well fires during the war. Additionally, after seeing 16 Gulf War veteran females in his gynecology clinic for the DoD registry, Wittich (32) concluded that a high proportion had gynecological problems during and after service in the Gulf.

#### Hospitalizations Studies

In-theater hospitalizations. Data are sparse regarding in-theater Gulf War hospitalizations. Electronic records were not routinely kept and some inpatient medical records have only been recently located. Wintermey and colleagues reviewed 574 admissions among US servicemen and Iraqi prisoners at an Army support hospital during the War. They concluded that hospitalization rates fluctuated with combat and that care to both US soldiers and Iraqi prisoners was of similar quality. Burkle et al. reviewed activities of two military field trauma centers located in Kuwait during the ground war and found that soldiers injured by land mines were most in need of emergency resuscitation. Recently, the Office of the Special Assistant for Gulf War

emergency resuscitation. (35) Recently, the Office of the Special Assistant for Gulf War Illnesses has made efforts to aggregate in-theater hospitalization records and electronically code admission data so that more detailed assessments may be possible.

Postwar hospitalization studies of electronic data. A number of controlled epidemiological studies of DoD hospitalizations have aided veterans and public health policymakers in examining evidence for increased postwar morbidity among Gulf War veterans. The first such study, published in 1996, compared Gulf War veterans and nondeployed veterans for postwar hospitalizations during 25 months after the war. (36) Adjusting for a number of covariates, including prewar hospitalization experience, Gulf War veterans who remained on active-duty were found to have increased postwar odds of hospitalizations in 5 of 45 multivariate statistical models. However, these differences were not consistent over the 25 months of study and were likely due to deferred medical care or other well understood postwar conditions.

DoD hospitalizations have also been studied for evidence that Gulf War veterans may be more prone than their non-deployed peers to suffer certain postwar diagnoses. Knoke et al. (37) examined the postwar diagnoses of the most common cancer among young males, testicular cancer, and found no evidence of increased risk among Gulf War veterans. In another study (38) the same team of investigators performed an epidemiological comparison to compile evidence of hospitalizations due to a new or as yet unidentified condition. Using an aggregate of 77 ICD-9 diagnoses previously identified by the Centers for Disease Control and Prevention in monitoring for unexplained deaths, they again compared regular active-duty Gulf War veteran and nondeployed veteran hospitalizations in DoD facilities. During the 50-month postwar period they found an increased hospitalization risk among Gulf War veterans beginning late in 1994. However, upon closer scrutiny, this increase was noted to be artificial and due to hospitalizations for non-medical reasons in support of the DoD's Gulf War veteran registry program in which veterans were admitted as inpatients in order to complete their diagnostic evaluation, not because they were ill.

Similarly, DoD hospitalization data were examined for evidence of unusual mental health morbidity that might be associated with the war. Dlugosz et al. (39) examined 30,539 initial mental health hospitalizations from June 1991 through September 1993 and found that Gulf War veterans were more likely to be admitted for alcohol-related disorders, drug-related disorders, and acute reactions to stress than their non-deployed peers. The strongest predictor of postwar mental health hospitalization was prewar mental health hospitalization. With the exception of alcohol-related disorders, the authors found no association between service in combat units during the Gulf War and risk of postwar mental health hospitalization. Rothberg et al. similarly found moderate but transient increases of postwar ambulatory mental health or social problems among Gulf War veterans at one medical treatment facility in the Southeastern United States. (40)

Smith et al.<sup>(41)</sup> compared Gulf War veterans with non-deployed veterans for the onset of three postwar conditions: systemic lupus erythematosus, amyotrophic lateral sclerosis, and fibromyalgia. Examining DoD hospital data from October 1988 through July 1997, they found no evidence that Gulf War veterans were more likely than their peers to be hospitalized for any of the three conditions.

#### **MENTION ALS STUDY**

Most recently, Blood et al. (42) compare the postwar hospitalization experience of US Marine deployed to the Gulf War with similar Marines deployed to the Vietnam conflict. They found similar hospitalization rates for both cohorts and no unexplained differences in the proportional distribution of diagnostic categories.

Because the hospitalization studies previously mentioned captured only data from personnel who were eligible for DoD hospital care, a comparison of Gulf War veteran and nondeployed veteran hospitalizations was published in 1998 using DoD, VA, and California nonfederal hospitalization data. (43) These analyses captured data from Reserve, and National Guard personnel, as well as former military personnel who no longer were eligible for federal healthcare. Because investigators were unable to determine accurate denominators for the VA and California hospital systems, they instead relied upon proportional morbidity ratios in making comparisons. There was no evidence that Gulf War veterans were suffering increased hospitalizations for infectious diseases, cancers, endocrine diseases, nervous system diseases, blood diseases, circulatory system diseases, musculoskeletal diseases, or skin conditions. Gulf War veterans had proportionally more hospitalizations for fractures, and bone and soft tissue conditions in DoD and California hospitals, and for respiratory diseases, digestive diseases, and vague symptom diagnoses in VA hospitals. concluded that while these findings are biologically plausible, they may simply be the result of as yet unidentified confounding risk factors, and they should be validated with further studies.

#### Postwar hospitalizations studies of cross-sectional survey data.

Postal surveys of military and former military personnel are another approach to comparing hospital morbidity among Gulf War veterans and their nondeployed peers. However, this method is more prone to several forms of bias. Kang et al. (44) reported that among the 11,441 Gulf War veterans and 9,476 nondeployed veterans they surveyed in 1996 and 1997, the proportion reporting a hospitalization due to illness during the last year was higher among Gulf War veterans (rate ratio = 1.22; 95% confidence interval (CI) 1.10 to 1.34). In another large postal survey conducted during 1997 through 1999, 12,049 Seabees (3,831 Gulf War veterans) who served during the Gulf War period were asked to complete questionnaires regarding their hospitalizations since 1990. Thirty-eight percent of Gulf War veterans and thirty-five percent of non-deployed veterans reported one or more hospitalizations.

#### Studies of Gulf War exposures and postwar hospitalizations.

DoD hospitalizations were additionally studied for evidence of Gulf War-related morbidity from possible chemical warfare exposures. Geographic and weather models were used to identify US Army soldiers possibly exposed to a nerve agent when a cache of nerve-agent laden rockets were destroyed after the war near Khamisiyah, Iraq, in March 1991. Although no evidence of acute nerve agent toxicity was documented, concern was raised regarding possible latent morbidity from subclinical nerve agent exposure. Using multivariate models, the postwar hospitalization experience of 348,291 US Army Gulf War veterans was studied through September 30, 1995. Possibly exposed Gulf War veterans were not found to be at increased risk of hospitalization for any cause, for diagnoses in any one of 15 ICD-9 large categories, or for diagnoses thought most likely to be associated with latent manifestations of subclinical nerve agent exposure.

#### OIL WELL HERE

Hospitalization studies, while not yielding evidence of a Gulf War syndrome or unusual Gulf War-related diagnoses, have demonstrated some increased risk of hospitalization among Gulf War veterans, particularly for mental health diagnoses and injuries. Data have also revealed that the most important predictors of postwar hospitalization are female gender, prewar hospitalization, healthcare occupation, enlisted rank, service in the Army, and low rank. (36, 38, 46) These studies have revealed the importance of considering numerous covariates when conducting future post-deployment research.

#### **OUTPATIENT VISITS**

#### In-theater outpatient visits.

A number of self-administered questionnaire surveys of US military personnel serving in the Gulf War theater were conducted before the fighting began. Richards et al. (47) surveyed 2,598 US military combat personnel serving in Saudi Arabia during September through December 1990 and found that respiratory disease symptoms were a frequent complaint. Among respondents, 43.1% complained of cough, 34.4% reported a sore throat, and 15.4% complained of chronic rhinorrhea. Hyams and colleagues (48) surveyed 2,022 soldiers stationed in various regions of Saudi Arabia during late 1,990. They found that with an average stay of only 2 months, 57% had at least one incident of diarrhea and 20% reported that diarrhea kept them from conducting their duties. Respiratory disease and diarrhea disease were similar common problems among deployed shipboard personnel. (49) The impact of infectious diseases upon Gulf War veterans is well described in two thorough reviews. (50, 51)

#### Postwar outpatient visits.

Relatively few controlled studies of postwar ambulatory care visits have been conducted among Gulf War veterans. This is due mainly to the lack of useful electronic outpatient records. The VA has electronic outpatient data (Outpatient Care File) available back to 1980, but the data lacked key epidemiological elements until improvements were made in 1996. (52) Similarly, outpatient electronic data (Standard Ambulatory Data Record) first became available for DoD health care facilities in 1996. Neither of these systems has been evaluated for use among Gulf War veterans and their peers.

A number of healthcare utilization reports are available from the Gulf War deployment period. Most are based on unofficial records and none include non-deployed controls. Hines<sup>(53)</sup> studied 10,165 ambulatory visits from an armor division during the Gulf War. He concluded that while women occasionally have needs that merit the care of a gynecologist, more broadly trained clinicians were usually well equipped to provide female soldier care. In a 1997 review article, Murphy et al.<sup>(54)</sup> came to similar conclusions but recommended better studies of gender-specific morbidity.

Hines<sup>(55)</sup> studied the 15,401 ambulatory visits made by Army personnel during 6 months of the Gulf War. He empirically created seven morbidity categories and compared male and female soldiers. He found that men were more likely to be seen for orthopedic or dermatological conditions, and women were more likely to be evaluated for psychiatric and optometric problems. Considering all visits of both male and female soldiers, ambulatory visits were most common for orthopedic problems (30.3%), respiratory problems (24.9%), and dermatological problems (13.9%). In a review of 6,772 visits among an US Army armored cavalry unit, Wasserman et al. (56) similarly found a high proportion of orthopedic, gastrointestinal, and respiratory problems among ambulatory visits during the Gulf War. Shaw et al. (57) found that among 1,820 medical encounters at two deployed Navy Fleet Hospitals, most frequent were injury category diagnoses (24%) and respiratory disease diagnoses (19%). Perhaps the most comprehensive in-theater ambulatory care evaluation was performed by Hyams, et al. (51) in conducting surveillance of ambulatory visits among 40,000 Marines stationed in northeastern Saudi Arabia. They found that gastroenteritis and acute respiratory track infections were most common. A gastroenteritis outbreak was abated after local fruits and vegetables were banned. A key element of success in determining the cause of diarrheal disease in the Gulf War theater was the availability of advance laboratory support. (58) They also noted that respiratory disease outbreaks occurred soon after large groups of deployed personnel arrived in theater.

#### Outpatient visits as captured by cross-sectional survey.

Another approach to evaluate ambulatory morbidity among Gulf War veterans has been to perform surveys of postwar healthcare utilization. In 1995, Stretch et al. (59)

reported a postal survey of 4,334 Gulf War veterans and their non-deployed peers finding that both active-duty and Reserve Gulf War veterans reported a higher prevalence than their non-deployed peers of visiting a physician within two weeks of completing the survey.

Similarly, in a 1998 report, (60) 3,113 Canadian Gulf War veterans and 3,439 non-deployed veterans were surveyed for contacts with healthcare professionals, hospital emergency room visits, and hospital admissions during the previous 12 months. The investigators found no evidence for increased healthcare visits but Gulf War veterans did report more nonprescription drug use and more medical conditions. Kang et al. (44) reported in their survey of more than 20,000 US Gulf War era service personnel that the Gulf War veterans reported more outpatient and inpatient visits than did their peers and they had more functional impairment, symptoms, and self-reported medical conditions.

#### **URANIUM EXPOSURE HERE**

While available data are not without problems, in general outpatient data suggest that Gulf War veterans are seeking healthcare more often than their non-deployed peers.

#### **ONGOING RESEARCH**

A number of studies are in progress to further examine healthcare utilization and diagnoses among Gulf War veterans. In a DoD and VA collaborative study, data from more than 120,000 participants in the two Gulf War veteran clinical registries will be analyzed for a more comprehensive understanding of the spectrum of illnesses among Gulf War veterans. In addition, the Uniformed Services University of Health Sciences (Bethesda, MD) is collaborating with the King Fahad National Guard Hospital (Riyadh Saudi Arabia), Naval Health Research Center (San Diego, CA), Centers for Disease Control and Prevention (Atlanta, GA), and DoD's Office of the Special Assistant for Gulf War Illnesses (Falls Church, VA) in examining the postwar healthcare utilization of Saudi National Guard Gulf War veterans compared with their Saudi National Guard colleagues who did not see combat. Finally, Naval Health Research Center, in collaboration with multiple DoD organizations, is comparing post-deployment morbidity between US veterans of the Gulf War and veterans of peacekeeping efforts in Southwest Asia and Bosnia.

#### DISCUSSION

Data from the Gulf War veteran health registries are influenced by self-selection bias and, as a result, are not representative of the entire Gulf War veteran population. Similarly, electronic hospitalization data were designed for administrative purposes and their use in epidemiological studies is not without problems. (61) Finally, self-reported data is understood to have many potential biases. McCauley et al. (62) found poor

validity and reliability in self-reported Gulf War exposures. Thus the reports we have summarized have considerable limitations but considered together lead us to a number of conclusions.

To date, the health registries and healthcare related epidemiological studies have failed to identify a unique constellation of symptoms or clinical signs that could be termed a "Gulf War syndrome." Instead, they demonstrated that the illnesses Gulf War veterans were suffering were similar to those experienced by other veterans, both historically and at present. (63) These studies have contributed much to reduce the anxiety related to alleged Gulf War related morbidity (Figure 1). Today, most studies have shown that Gulf War veterans are not suffering unexplained mortality, cancers, unusual hospitalizations or from chronic diseases any more than their non-deployed Most researchers currently recognize that the conditions most strongly associated with Gulf War service are those involving unexplained symptoms. Hence, the focus on present and future medical research among Gulf War veterans and other military populations has shifted to testing empirical treatments (64,65) proved effective among other populations suffering from symptom-based conditions and to evaluating how the health of Gulf War veterans changes over time. (66) A large population-based study<sup>(67)</sup> is being planned to follow the health of Gulf War veterans and other US military personnel over the next 20 years.

#### **CONCLUSIONS AND FUTURE RESEARCH NEEDS**

Gulf War veterans have suffered a wide range of health problems since returning from the war. However, health care registry data and epidemiological studies of health-care utilization have revealed no unique Gulf War syndrome. Instead, they have demonstrated conditions similar to those affecting other military and adult populations, often with no identified increase in risk. However, regarding multi-symptom conditions, Gulf War veterans do seem to be suffering increased morbidity and to be using healthcare services more than their peers. Such multi-symptom conditions are now the focus of significant research.

In addition to evaluating possible therapies for multi-symptom conditions, federal research should now focus on identifying risk factors for the development of post-deployment symptom-based conditions. In fact, plans are in progress to evaluate pre-deployment screening strategies to identify new military members at greatest risk of post-deployment symptom-based illness. (68) Such risk factor identification is necessary to guide future preventive efforts to reduce post-deployment morbidity.

#### **ACKNOWLEDGEMENTS**

Dr. Gray was formerly Director of the Center for Deployment Health Research, at Naval Health Research Center, San Diego, CA; which an early draft (dated 7-31-00) of this manuscript was prepared.

#### REFERENCES

- 1. Nelson SS (1992). Sick Gulf vets demand care for mystery illness. <u>Navy Times</u>, 21.
- 2. Beale P (1994). Gulf illness. BMJ, 308:1574.
- 3. Mystery illness and the Gulf War. MacLean's, 1993.
- 4. DeFraites RF, Wanat ER, Norwood AE, Williams S, Cowan D, Callahan T. (1992)

  Investigation of a Suspected Outbreak of an Unknown Disease Among Veterans of
  Operation Desert Shield/Storm, 123rd Army Reserve Command, Fort Benjamin
  Harrison, Indiana, April 1992. Washington, DC: Epidemiology Consultant Service,
  Division of Preventive Medicine, Walter Reed Army Institute of Research.
- Institute of Medicine (1995). <u>Health Consequences of Service During the Persian</u>
   <u>Gulf War: Initial Findings and Recommendations for Immediate Action</u>.
   Washington, DC: National Academy Press.
- 6. Murphy FM (1999). Gulf war syndrome. <u>BMJ</u>, 318:274-5.
- 7. Joseph S (1997), The Comprehensive Clinical Evaluation Program Evaluation Team. A comprehensive clinical evaluation of 20,000 Persian Gulf War veterans. Mil Med, 162:149-155.
- 8. Coker WJ, Bhatt BM, Blatchley NF, Graham JT (1999). Clinical findings for the first 1000 Gulf war veterans in the Ministry of Defence's medical assessment programme. <u>BMJ</u>, 318:290-4.
- 9. Robinson A (1995). Veterans worry that unexplained medical problems a legacy of service during Gulf War. <u>CMAJ</u>,152:944-7.
- Department of Veterans Affairs. <u>Annual Report to Congress: Federally Sponsored Research on Persian Gulf Veterans' Illnesses for 1998</u>. Washington, DC: Department of Veterans Affairs, The Research Working Group of the Persian Gulf Veterans Coordinating Board, 1999.
- 11. Riddle JR, Hyams KC, Murphy FM, Mazzuchi JF (2000). In the borderland between health and disease following the Gulf War. Mayo Clin Proc. 75:777-9.
- 12. Kang H, Bullman T, MacFarlane G, Gray G. Mortality among US and UK veterans of the Persian Gulf War: A review. <u>J Occup Environ Med</u> 2002 (in press).
- 13. Barrett D, Gray G, Doebbeling B, Clauw D, Reeves W, Board' t PGVC. The prevalence of symptoms and symptom-based conditions among Gulf War veterans: current status of research findings. <u>Epidemiol Rev</u> 2002 (under review).
- 14. Hyams KC, Riddle J, Trump DH, Graham JT (2001). Endemic infectious diseases and biological warfare during the Gulf War: a decade of analysis and final concerns. Am J Trop Med Hyg, 65:664-70.

- 15. Brown MA, Murphy FM, Mather SH (2002). Innovation in veterans' health care and assistance: the Department of Veterans Affairs 10 years after the Gulf War. Mil Med,167:191-5.
- 16. Trump DH, Mazzuchi JF, Riddle J, Hyams KC, Balough B (2002). Force health protection: 10 years of lessons learned by the Department of Defense. Mil Med, 167:179-85.
- 17. Henry L, Gray G (1999). Topical Bibliography of Published Works Regarding the Health of Veterans of the Persian Gulf War. Technical Document 99-3C, San Diego: Naval Health Research Center.
- Gray GC, Hawksworth AW, Smith TC, Kang HK, Knoke JD, Gackstetter GD (1998). Gulf War Veterans' Health Registries. Who is most likely to seek evaluation? <u>Am J Epidemiol</u>, 148:343-9.
- 19. Murphy FM, Kang H, Dalager NA, et al (1999). The health status of Gulf War veterans: lessons learned from the Department of Veterans Affairs Health Registry. Mil Med, 164:327-31.
- 20. Smith T, Smith B, Ryan M, et al. Ten years and 100,000 participants later:

  Occupational factors influencing enrollment in US Gulf War clinical registries. J.

  Occup Environ Med 2002 (in press).
- 21. Lee HA, Gabriel R, Bale AJ, Bolton P, Blatchley NF (2001). Clinical findings of the second 1000 UK GulfWar veterans who attended the Ministry of Defence's Medical Assessment Programme. J R Army Med Corps, 147:153-60.
- 22. Stuart JA, Murray KM, Ursano RJ, Wright KM (2002). The Department of Defense's Persian Gulf War registry year 2000: an examination of veterans' health status. Mil Med,167:121-8.
- 23. Kroenke K, Koslowe P, Roy M (1998). Symptoms in 18,495 Persian Gulf War veterans. <u>J Occup Environ Med</u>, 40:520-528.
- 24. Escalante A, Fischbach M (1998). Musculoskeletal manifestations, pain, and quality of life in Persian Gulf War veterans referred for rheumatologic evaluation. <u>J</u> Rheumatol, 25:2228-35.
- 25. Grady EP, Carpenter MT, Koenig CD, Older SA, Battafarano DF (1998). Rheumatic findings in Gulf War veterans. <u>Arch Intern Med</u>, 158:367-371.
- 26. Erickson AR, Enzenauer RJ, Bray VJ, West SG (1998). Musculoskeletal complaints in Persian Gulf War veterans. Journal of Clinical Rheumatology, 4:181-185.
- 27. Newmark J, Clayton WLI (1995). Persian Gulf illnesses: preliminary neurological impressions. Mil Med, 160:505-507.
- 28. Roy M, Koslowe P, Kroenke K, Magruder C (1998). Signs, symptoms, and ill-defined conditions in Persian Gulf War veterans: findings from the Comprehensive Clinical Evaluation Program. <u>Psychosom Med</u>, 60:663-668.

- 29. The International Classification of Diseases, 9th Revision, Clinical Modification. 3rd ed. Washington, DC: US Department of Health and Human Services, 1991.
- 30. Kipen HM, Hallman W, Kang H, Fiedler N, Natelson BH (1999). Prevalence of chronic fatigue and chemical sensitivities in Gulf Registry Veterans. <u>Arch Environ Health</u>, 54:313-8.
- 31. Das AK, Davanzo LD, Poiani GJ, et al. (1999). Variable extrathoracic airflow obstruction and chronic laryngotracheitis in Gulf War veterans. Chest, 115:97-101.
- 32. Wittich AC (1996). Gynecologic evaluation of the first female soldiers enrolled in the Gulf War Comprehensive Clinical Evaluation Program at Tripler Army Medical Center. Mil Med,161:635-637.
- Wintermeyer SF, Pina JS, Cremins JE, Heier JS (1994). The inpatient experience of a US Army combat support hospital in the Persian Gulf during non-combat and combat periods. <u>Mil Med</u>, 159:746-51.
- 34. Wintermeyer SF, Cremins JE, Pina JS, Heier JS (1996). Medical care of Iraqis at a forwardly deployed US Army hospital during Operation Desert Storm. Mil Med, 161:294-297.
- 35. Burkle Jr FM, Newland C, Meister SJ, Blood CG (1994). Emergency medicine in the Persian Gulf War--Part 3: battlefield casualties. <u>Ann Emerg Med</u>, 23:755-760.
- 36. Gray GC, Coate BD, Anderson CM, et al. (1996). The postwar hospitalization experience of U.S. veterans of the Persian Gulf War. N Engl J Med. 335:1505-13.
- 37. Knoke JD, Gray GC, Garland FC (1998). Testicular cancer and Persian Gulf War service. <u>Epidemiology</u>, 9:648-653.
- 38. Knoke JD, Gray GC (1998). Hospitalizations for unexplained illnesses among U.S. veterans of the Persian Gulf War. <u>Emerg Infect Dis</u>, 4:211-9.
- 39. Dlugosz LJ, Hocter WJ, Kaiser KS, et al. (1999). Risk factors for mental disorder hospitalization after the Persian Gulf War: U.S. Armed Forces, June 1, 1991-September 30, 1993. <u>J Clin Epidemiol</u>, 52:1267-1278.
- 40. Rothberg JM, Koshes RJ, Shanahan J, Christman K (1994). Desert Shield deployment and social problems on a US Army combat support post. Mil Med, 159:246-248.
- 41. Smith TC, Gray GC, Knoke JD (2000). Is systemic lupus erythematosus, amyotrophic lateral sclerosis, or fibromyalgia associated with Persian Gulf War service? An examination of Department of Defense hospitalization data. <u>Am J Epidemiol</u>, 51:1053-9.
- 42. Blood CG, Aboumrad TL (2001). A comparison of postdeployment hospitalization incidence between active duty Vietnam and Persian Gulf War Veterans. Mil Med, 166:648-55.

- 43. Gray GC, Smith TC, Kang HK, Knoke JD (2000). Are Gulf War veterans suffering war-related illnesses? Federal and civilian hospitalizations examined, June 1991 to December 1994. Am J Epidemiol,151:63-71.
- 44. Kang HK, Mahan CM, Lee KY, Magee CA, Murphy FM (2000). Illnesses among United States veterans of the Gulf War: a population-based survey of 30,000 veterans. J Occup Environ Med, 42:491-501.
- 45. Kaiser K, Reed R, Gastanaga V, Gray G. An analysis of morbidity with unclear etiology in a population of US Navy Gulf War veterans. Conference on Federally Sponsored Gulf War Veterans' Illnesses Research. June 23-25, Crystal City, Virginia, 1999.
- 46. Gray GC, Smith TC, Knoke JD, Heller JM (1999). The postwar hospitalization experience of Gulf War veterans possibly exposed to chemical munitions destruction at Khamisiyah, Iraq. <u>Am J Epidemiol</u>,150:532-40.
- 47. Richards AL, Hyams KC, Watts DM, Rozmajzl PJ, Woody JN, Merrell BR (1993). Respiratory disease among military personnel in Saudi Arabia during Operation Desert Shield. Am J Public Health, 83:1326-9.
- 48. Hyams KC, Bourgeois AL, Merrell BR, et al. (1991). Diarrheal disease during Operation Desert Shield [see comments]. N Engl J Med, 325:1423-8.
- 49. Paparello SF, Garst P, Bourgeois AL, Hyams KC (1993). Diarrheal and respiratory disease aboard the hospital ship, USNS-Mercy T-AH 19, during Operation Desert Shield. Mil Med,158:392-5.
- 50. Hyams K (1999). Gulf War syndrome: potential role of infectious diseases. <u>Current Opinion in Infectious Diseases</u>,12:439-443.
- Hyams KC, Hanson K, Wignall FS, Escamilla J, Oldfiled EC (1995). The impact of infectious diseases on the health of U.S. troops deployed to the Persian Gulf during operations Desert Shield and Desert Storm. <u>Clin Infect Dis</u>, 20:1497-1504.
- 52. Boyko EJ, Koepsell TD, Gaziano JM, Horner RD, Feussner JRm (2000). US Department of Veterans Affairs medical care system as a resource to epidemiologists. <u>Am J Epidemiol</u>,151:307-314.
- 53. Hines JF (1992). Ambulatory health care needs of women deployed with a heavy armor division during the Persian Gulf War. Mil Med, 57:219-21.
- 54. Murphy F, Browne D, Mather S, Scheele H, Hyams KC (1997). Women in the Persian Gulf War: health care implications for active duty troops and veterans. Mil Med, 162:656-660.
- 55. Hines JF (1993). A comparison of clinical diagnoses among male and female soldiers deployed during the Persian Gulf War. Mil Med, 158:99-101.
- 56. Wasserman GM, Martin BL, Hyams KC, Merrill BR, Oaks HG, McAdoo HA (1997). A survey of outpatient visits in a United States Army Forward Unit during Operation Desert Shield. <u>Mil Med</u>, 162:374-379.

- 57. Shaw E, Hermansen L, Pugh W, White M (1991). Disease and non-battle injuries among Navy and Marine Corps personnel during operation Desert Shield /Desert Storm. Report 91-44. San Diego: Naval Health Research Center.
- 58. Hyams KC, Bourgeois AL, Escamilla J, Burans J, Woody JN (1993). The Navy forward laboratory during Operations Desert Shield/Desert Storm. Mil Med, 158:729-732.
- 59. Stretch RH, Bliese PD, Marlowe DH, Wright KM, Knudson KH, Hoover CH (1995). Physical health symptomatology of Gulf War-era service personnel from the states of Pennsylvania and Hawaii. Mil Med, 160:131-136.
- 60. Goss Gilroy Inc. Health study of Canadian forces personnel involved in the 1991 conflict in the Persian Gulf Vol I. Ottawa, Ontario, 1998.
- 61. Kuller L (1995). The use of existing databases in morbidity and mortality studies. Am J Public Health, 85:1198-1200.
- 62. McCauley LA, Joos SK, Spencer PS, Lasarev M, Shuell T (1999). Strategies to assess validity of self-reported exposures during the Persian Gulf War. Portland Environmental Hazards Research Center. Environ Res. 81:195-205.
- 63. Hyams KC, Wignall FS, Roswell R (1996). War syndromes and their evaluation: from the US Civil War to the Persian Gulf War. <u>Ann Intern Med</u>,125:398-405.
- 64. Guarino P, Peduzzi P, Donta ST, et al. (2001). A multicenter two by two factorial trial of cognitive behavioral therapy and aerobic exercise for Gulf War veterans' illnesses: design of a veterans affairs cooperative study (CSP #470). Control Clin Trials, 22:310-32.
- 65. Gerrity TR, Feussner JR (1999). Emerging research on the treatment of Gulf War veterans' illnesses. J Occup Environ Med, 41:440-2.
- 66. Institute of Medicine: Committee on Measuring the Health of Gulf War Veterans. Gulf War Veterans: Measuring Health. Washington, DC: National Academy Press, 1999.
- 67. Chesbrough K, Ryan M, Amoroso A, et al. The Millennium Cohort Study: A 21-year prospective cohort study of 140,000 military personnel. Mil Med 2002 (in press).
- 68. Hyams KC, Barrett DH, Duque D, et al. (2002). The Recruit assessment Program: a program to collect comprehensive baseline health data from U.S. military personnel. Mil Med,167:44-7.

Table 1. Special Gulf War veteran health evaluations.

Sponsor	Registry Name	Start Date	Number of Veterans Evaluated as of June 2000	Number Deployed	
United States Department of Veterans Affairs	Persian Gulf Veterans Health Registry (6)	August 1992	70,000+	697,000	
United States Department of Defense	Comprehensive Clinical Evaluation Program (7)	June 1994	53,000+	337,000	
United Kingdom Ministry of Defence	Gulf War Medical Assessment Programme (8)	July 1993	3,000+	53,000	
Canada Department of National Defence	Canadian Gulf War Registry (9)	January 1995	226+	4,500	

Table 2. Prevalence of frequently reported symptoms, by Gulf War veteran registry.

		· · · · · · · · · · · · · · · · · · ·		
	US	÷		
	Department of	US		
	Veterans	Department of	United	
	Affairs (19)	Defense	Kingdom	Canada
÷	(n = 52,835)	( <i>n</i> = 53,032)*	(n = 1000)(8)	$(n = 104)\dagger$
Symptom	Symptom prevalence (%)			
Fatigue	20.5	37.2	42.1	86.5
Skin rash	18.4	24.0	19.4	47.1
Headache	18.0	32.7	25.6	65.4
Muscle and	16.8	40.9‡	39.5	75.0
joint pain	10.0	40.94	39.5	70.0
Cognitive	14.0	33.5**	26.1	84.6
problems	14.0	00.0	20.1	•
Shortness of	7.9	19.0	24.3	33.0
breath	7.5	10.0	24.0	
Sleep	5.9	29.1 <sup>††</sup>	21.2	74.0
disturbance	0.5	20.1	- 1	74.0
Diarrhea and				
other	4.6	27.0 <sup>‡‡</sup>	21.8	55.8
gastrointestinal				
No complaint	12.3	41.1***	7.4	0

<sup>\*</sup>As of May 31, 2000.

<sup>&</sup>lt;sup>†</sup> From 104 veterans evaluated at Ottawa's Gulf War Clinic April 1995 to December 1997.

<sup>&</sup>lt;sup>‡</sup>Answered "yes" to "muscle pain," "joint pain," or both.

<sup>\*\*</sup> Answered "yes" to "difficulty concentrating," "memory loss," or both.

<sup>&</sup>lt;sup>††</sup> Answered "yes" to any of the following: "sleep disturbance," "loud snoring," or "stopped breathing while sleeping."

<sup>&</sup>lt;sup>‡‡</sup> Answered "yes" to "abdominal pain", "diarrhea", or both.

<sup>\*\*\*</sup>Did not answer "yes" to 18 questions about symptoms including "Do you have any other symptoms?"

Table 3. Percent distribution of selected diagnoses for Gulf War veterans participating in a health registry

		US		
		Department	US	
,		of Veterans	Department	
		Affairs*	of Defense <sup>†</sup>	
Diagnoses		(n = 69,941)	(n = 53,032)	
Diagnoses ICD-9 code <sup>‡</sup>		% of diagnoses		
None	**	27.0	29.5	
Malaise and	780.7	22.0	8.4	
fatigue	700.7	22.0		
Headache	784.0	21.0	6.2	
Pain in joint	719.4	13.2	12.7	
Sleep disturbances	780.5	7.9	7.9	
Depressive disorder, not elsewhere classified	311	4.1	4.9	
Lumbago	724.2	4.0	6.1	
Psychalgia	307.8	3.9	7.7	
Other specified adjustment reactions	309.8	3.9	3.6	
Essential hypertension, unspecified	401.9	3.6	2.8	
Contact dermatitis and other eczema, unspecified cause	692.9	3.6	2.6	
Unspecified sinusitis (chronic)	473.9	3.2	1.9	

Other and unspecified noninfectious gastroenteritis and colitis	558.9	2.8	1.5
Asthma, unspecified	493.9	2.7	2.5
Migraine, unspecified	346.9	2.6	3.8
Allergic rhinitis, cause unspecified	477.9	2.5	3.4
Anxiety states	300.0	2.4	1.7
Osteoarthrosis, unspecified	715.9	2.2	4.4
Alopecia	704.0	2.1	2.4
Irritable colon	564.1	1.9	3.7

<sup>\*</sup>In the VA registry a special non-ICD-9 code "DX111" was used as well as ICD-9 code V65.5 (person with feared complaint in whom no diagnoses were made) to indicate veterans with no diagnoses. All are considered as missing.

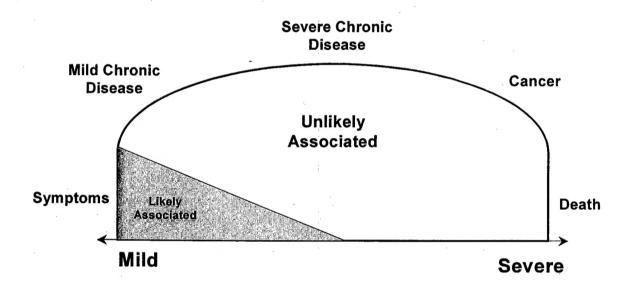
<sup>&</sup>lt;sup>†</sup> Percentage of 69,941 Gulf War veterans who completed the VA registry examination as of September 30, 1999.

<sup>&</sup>lt;sup>‡</sup> 4-digit code from *International Classification of Diseases, Ninth Revision, Clinical Modification, 3rd Edition* (29).

Data as of May 30, 2000 – Diagnoses were reported in the any of the primary or 6 secondary diagnostic fields. A code of "xxx.xx" was interpreted as missing.

#### **LEGEND**

Figure 1. Health registry studies and epidemiological studies have demonstrated no unique Gulf War syndrome and no increased risk among Gulf War veterans of unexplained mortality, cancer, severe chronic disease, or mild chronic disease. However, it is apparent that Gulf War veterans are suffering from increased symptoms and this spectrum of morbidity is now the focus of much federal research.



REPORT DOCUMENTATION PAGE	
The public reporting burden for this collection of information is estimated to average 1 hour per response, including sources, gathering and maintaining the data needed, and completing and reviewing the collection of information. S aspect of this collection of information, including suggestions for reducing the burden, to Washington Headquarters 1215 Jefferson Davis Highway, Suite 1204, Arlington, VA 22202-4302, Respondents should be aware that notwiths to any penalty for failing to comply with a collection of information if it does not display a currently valid OMB Contro THE ABOVE ADDRESS.	end comments regarding this burden estimate or any other Services, Directorate for Information Operations and Reports, tanding any other provision of law, no person shall be subject I number. PLEASE DO NOT RETURN YOUR FORM TO
1. Report Date (DD MM YY)  2. Report Type  13 Jul 2000  New	3. DATES COVERED (from - to) 1996 to Feb 200
4. TITLE AND SUBTITLE Gulf War Veteran Healthcare Utilization: After more than 10 years of study, what have we learned?  6. AUTHORS Gregory C. Gray, Gary D. Gackstetter, Han K. Kang, John T. Graham, & Ken C. Scott  7. PERFORMING ORGANIZATION NAME(S) AND ADDRESS(ES) Naval Health Research Center P.O. Box 85122	5a. Contract Number: 5b. Grant Number: 5c. Program Element: 63738D 5d. Project Number: P4464 5e. Task Number: 001 5f. Work Unit Number: 60002
San Diego, CA 92186-5122  8. SPONSORING/MONITORING AGENCY NAMES(S) AND ADDRESS(ES) Chief, Bureau of Medicine and Surgery	9. PERFORMING ORGANIZATION REPORT NUMBER Report 00-26
M2 2300 E St NW	10. Sponsor/Monitor's Acronyms(s) BUMED
Washington DC 20372-5300	11. Sponsor/Monitor's Report Number(s)
12 DISTRIBUTION/AVAILABILITY STATEMENT Approved for public release; distribution unlimited.	
13. SUPPLEMENTARY NOTES	
Purpose: To review clinical and epidemiological studies of morbidity among Gulf special healthcare registries, hospitalizations, and outpatient visits.  Data Sources: MEDLINE, Reports to Congress, and a topical Gulf War veteran tresearch, ongoing research, expert panel reviews, and case reports relevant to the Study Selection: Studies published in peer-reviewed medical journals.  Data Extraction: Published reports are reviewed and findings summarized.  Data Synthesis: Study approaches and findings are summarized and considered considered.  Conclusions: Published health registry studies and healthcare-related epidemiologic increased Gulf War veterans access to care, and helped to rule out severe warguided empirical therapies for the excessive number of Gulf War veterans suffer	bibliography were used to identify published he objective.  I in aggregate. Additional studies are object research have reduced speculation, elated morbidity. In addition, they have
15. SUBJECT TERMS  Veteran Healthcare Utilization, Gulf War Veterans, healthcare registries, hospit	
16. SECURITY CLASSIFICATION OF: 17. LIMITATION 18. NUMBER 19a. NAM	E OF RESPONSIBLE PERSON

OF ABSTRACT

UNCL

a. REPORT

UNCL

b.ABSTRACT | b. THIS PAGE

UNCL

UNCL

**OF PAGES** 

21

19b. TELEPHONE NUMBER (INCLUDING AREA CODE) COMM/DSN: (619) 553-8429

Commanding Officer